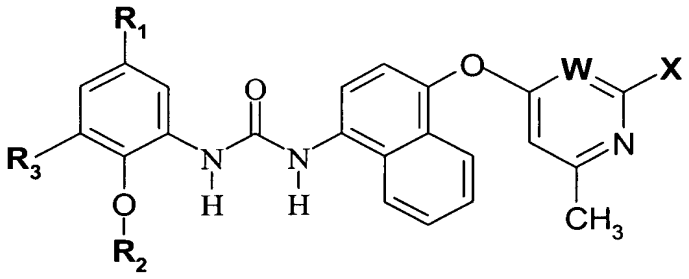
LISTING OF CLAIMS

Claim 1 (currently amended): A compound of the formula (I):



(I)

wherein

$\mathbf{R_1}$ is $-CF_3$, $-CH(CH_3)(CF_3)$, $-CH(CF_3)_2$, $-OCF_3$, $-CF_2CF_3$;

$\mathbf{R_2}$ is $C_{1\text{-}5}$ alkyl;

$\mathbf{R_3}$ is attached at the 3- or 4-position on the phenyl ring and is hydrogen, $-NH_2$ or $\mathbf{R_4}$-$S(O)_2$-NH- wherein $\mathbf{R_4}$ is chosen from $C_{1\text{-}5}$ alkyl or carbocycle;

W is ~~CH or~~ an N atom;

X is chosen from

C1-5 alkyl or C1-5 alkoxy each optionally substituted by mono- or di-C1-3 alkyl amino ~~,morpholinyl, piperazinyl, pyrrolidinyl, triazolyl, imidazolyl or piperadinyl each ring being further optionally substituted with C1-3 alkyl~~ ;

or **X** is $-N(\mathbf{R^a})_2$ wherein $\mathbf{R^a}$ is independently chosen from hydrogen, C1-5 alkyl, aryl, arylC1-3 alkyl, C3-7cycloalkyl, C3-7cycloalkyl C1-3 alkyl , and C1-5alkoxyC1-5alkyl ~~and heterocyclyl C1-3 alkyl wherein the heterocycyl is chosen from tetrahydrofuran,~~

~~pyrrolidinyl and morpholinyl,~~ each $\mathbf{R^a}$ where possible is optionally substituted by one to two C1-5 alkyl, C1-5 alkoxy, hydroxy, halogen or amino optionally mono- or di-substituted by C1-3 alkyl;

or the pharmaceutically acceptable acids, esters, salts or isomers thereof.

Claim 2 (currently amended): The compound according to claim 1 wherein:

$\mathbf{R_1}$ is $-CF_3$, $-CH(CH_3)(CF_3)$, $-CH(CF_3)_2$, $-OCF_3$ or $-CF_2CF_3$;

$\mathbf{R_2}$ is C_{1-3} alkyl;

~~W is an N atom;~~

X is chosen from
C1-5 alkyl or C1-5 alkoxy each optionally substituted by mono- or di-C1-3 alkyl amino ~~, morpholinyl, piperazinyl, pyrrolidinyl, triazolyl, imidazolyl or piperadinyl each ring being further optionally substituted with C1-3 alkyl~~ ;

or **X** is $-N(\mathbf{R^a})_2$ wherein $\mathbf{R^a}$ is independently chosen from hydrogen, C1-5 alkyl, phenylC1-3 alkyl, C3-6cycloalkyl, C3-6cycloalkyl C1-3 alkyl ~~,~~ and C1-3alkoxyC1-3alkyl ~~and heterocyclyl C1-3 alkyl wherein the heterocycyl is chosen from tetrahydrofuran, pyrrolidinyl and morpholinyl,~~ each $\mathbf{R^a}$ where possible is optionally substituted by one to two C1-3 alkyl, C1-3 alkoxy, hydroxy, halogen or amino optionally mono- or di-substituted by C1-2 alkyl.

Claim 3 (original): The compound according to claim 2 wherein:

$\mathbf{R_1}$ is $-CF_3$, $-CH(CH_3)(CF_3)$, $-CH(CF_3)_2$ or $-CF_2CF_3$;

R_2 is $C_{1\text{-}2}$ alkyl.

Claim 4(original): The compound according to claim 3 wherein:

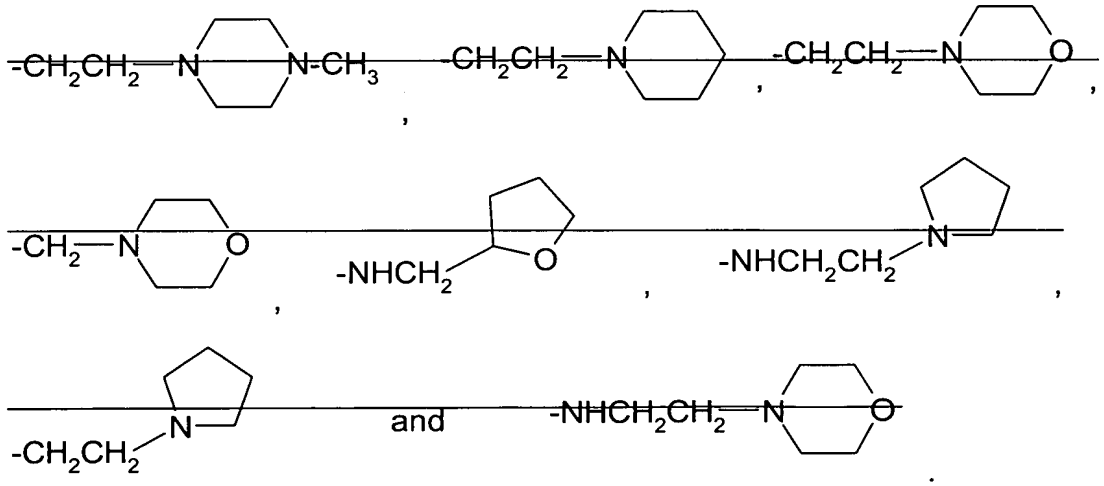
R_1 is $-CF_3$;

R_2 is $-CH_3$.

Claim 5 (currently amended): The compound according to claim 4 wherein:

X is chosen from

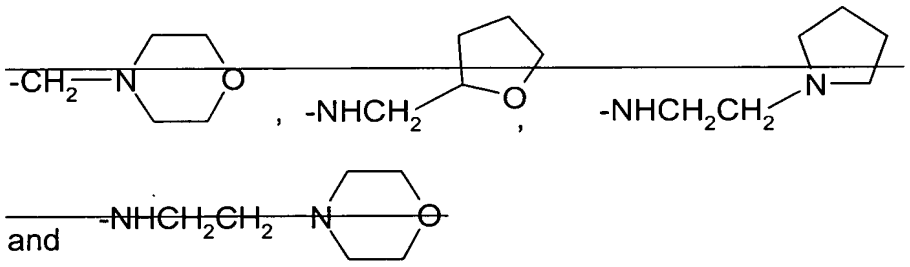
NH_2, $NH(CH_3)$, $-NHCH_2CH_2N(CH_3)_2$, $-NHCH_2CH_2\text{-}O\text{-}CH_3$, $-CH_2\text{-}N(CH_3)_2$, $-CH_2CH_2\text{-}N(CH_3)_2$, $-CH_2CH_2CH_2\text{-}N(CH_3)_2$, $-CH_2CH_2CH_2CH_2\text{-}N(CH_3)_2$, $-CH_2\text{-}NH(CH_3)$, $-CH_2CH_2\text{-}NH(CH_3)$, $-CH_2CH_2CH_2\text{-}NH(CH_3)$, $-CH_2CH_2CH_2CH_2\text{-}NH(CH_3)$, $-O\text{-}CH_2\text{-}N(CH_3)_2$, $-O\text{-}CH_2CH_2\text{-}N(CH_3)_2$, $-O\text{-}CH_2CH_2CH_2\text{-}N(CH_3)_2$, $-O\text{-}CH_2CH_2CH_2CH_2\text{-}N(CH_3)_2$, $-OCH_2\text{-}NH(CH_3)$, $-O\text{-}CH_2CH_2\text{-}NH(CH_3)$, $-O\text{-}CH_2CH_2CH_2\text{-}NH(CH_3)$, and $-O\text{-}CH_2CH_2CH_2CH_2\text{-}NH(CH_3)$,



Claim 6 (currently amended): The compound according to claim 5 wherein:

X is chosen from

NH_2, $NH(CH_3)$, $-NHCH_2CH_2N(CH_3)_2$, $-NHCH_2CH_2-O-CH_3$, $-CH_2-N(CH_3)_2$, $-CH_2CH_2-N(CH_3)_2$, $-CH_2CH_2CH_2-N(CH_3)_2$, $-CH_2CH_2CH_2CH_2-N(CH_3)_2$, $-CH_2-NH(CH_3)$, $-CH_2CH_2-NH(CH_3)$, $-CH_2CH_2CH_2-NH(CH_3)$, $-CH_2CH_2CH_2CH_2-NH(CH_3)$, $-O-CH_2-N(CH_3)_2$, $-O-CH_2CH_2-N(CH_3)_2$, $-O-CH_2CH_2CH_2-N(CH_3)_2$, $-O-CH_2CH_2CH_2CH_2-N(CH_3)_2$, $-OCH_2-NH(CH_3)$, $-O-CH_2CH_2-NH(CH_3)$, $-O-CH_2CH_2CH_2-NH(CH_3)$ ~~,~~ and $-O-CH_2CH_2CH_2CH_2-NH(CH_3)$ ~~,~~



Claim 7 (currently amended): A compound chosen from:

1-[4-(2-Amino-6-methyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea;
~~1-(2-Methoxy-5-trifluoromethyl-phenyl)-3-[4-(6-methyl-2-morpholin-4-ylmethyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-urea;~~
1-{4-[2-(2-Dimethylamino-ethylamino)-6-methyl-pyrimidin-4-yloxy]-naphthalen-1-yl}-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea;
1-{4-[2-(2-Methoxy-ethylamino)-6-methyl-pyrimidin-4-yloxy]-naphthalen-1-yl}-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea;

~~1-(2-Methoxy-5-trifluoromethyl-phenyl)-3-(4-{6-methyl-2-[(tetrahydro-furan-2-ylmethyl)-amino]-pyrimidin-4-yloxy}-naphthalen-1-yl)-urea;~~
~~1-(2-Methoxy-5-trifluoromethyl-phenyl)-3-{4-[6-methyl-2-(2-pyrrolidin-1-yl-ethylamino)-pyrimidin-4-yloxy]-naphthalen-1-yl}-urea;~~
~~1-(2-Methoxy-5-trifluoromethyl-phenyl)-3-{4-[6-methyl-2-(2-morpholin-4-yl-ethylamino)-pyrimidin-4-yloxy]-naphthalen-1-yl}-urea;~~
1-[4-(2-Cyclopropylamino-6-methyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea;
1-{4-[2-(2-Methoxy-1-methyl-ethylamino)-6-methyl-pyrimidin-4-yloxy]-naphthalen-1-yl}-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea;
1-{4-[2-(2,3-Dihydroxy-propylamino)-6-methyl-pyrimidin-4-yloxy]-naphthalen-1-yl}-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea;
1-[4-(2,6-Dimethyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea;
1-[4-(2-Methoxy-6-methyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea and
N-(3-{3-[4-(2,6-Dimethyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-ureido}-2-methoxy-5-trifluoromethyl-phenyl)-methanesulfonamide

or the pharmaceutically acceptable acids, esters, salts or isomers thereof.

Claim 8 (currently amended): A compound chosen from:

1-[4-(2-Ethylamino-6-methyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea;
1-{4-[2-(Cyclopropylmethyl-amino)-6-methyl-pyrimidin-4-yloxy]-naphthalen-1-yl}-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea;
1-[4-(2-Isopropylamino-6-methyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea;
1-[4-(2-Amino-6-methyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-3-[2-methoxy-5-(2,2,2-trifluoro-1-methyl-ethyl)-phenyl]-urea;
1-[4-(2-Amino-6-methyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-3-[2-methoxy-5-(2,2,2-trifluoro-1-trifluoromethyl-ethyl)-phenyl]-urea;
1-[4-(2-Amino-6-methyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-3-[2-methoxy-5-(2,2,2-trifluoro-ethyl)-phenyl]-urea;
1-[4-(2-Methylamino-6-methyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-3-(2-methoxy-5-trifluoromethoxy-phenyl)-urea;
1-(2-Methoxy-5-trifluoromethoxy-phenyl)-3-[4-(6-methyl-2-ethylamino-pyrimidin-4-yloxy)-naphthalen-1-yl]-urea;
1-{4-[2-(2-Dimethylamino-ethylamino)-6-methyl-pyrimidin-4-yloxy]-naphthalen-1-yl}-3-(2-methoxy-5-trifluoromethoxy-phenyl)-urea;
~~1-(2-Methoxy-5-trifluoromethoxy-phenyl)-3-{4-[6-methyl-2-(2-pyrrolidin-1-yl-ethylamino)-pyrimidin-4-yloxy]-naphthalen-1-yl}-urea;~~
1-{4-[2-(2-Methoxy-ethylamino)-6-methyl-pyrimidin-4-yloxy]-naphthalen-1-yl}-3-(2-methoxy-5-trifluoromethoxy-phenyl)-urea;
~~1-(2-Methoxy-5-trifluoromethoxy-phenyl)-3-[4-(6-methyl-2-morpholin-4-ylmethyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-urea;~~
~~1-(2-Methoxy-5-trifluoromethoxy-phenyl)-3-{4-[6-methyl-2-(4-methyl-piperazin-1-ylmethyl)-pyrimidin-4-yloxy]-naphthalen-1-yl}-urea;~~

~~1-(2-Methoxy-5-trifluoromethoxy-phenyl)-3-[4-(6-methyl-2-pyrrolidin-1-ylmethyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-urea;~~
1-[4-(2-Dimethylaminomethyl-6-methyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-3-(2-methoxy-5-trifluoromethoxy-phenyl)-urea;
~~1-{4-[2-(2-Imidazol-1-yl-ethoxy)-6-methyl-pyrimidin-4-yloxy]-naphthalen-1-yl}-3-(2-methoxy-5-trifluoromethoxy-phenyl)-urea;~~
~~1-[2-Methoxy-5-(2,2,2-trifluoro-ethyl)-phenyl]-3-{4-[6-methyl-2-(1-methyl-piperidin-4-ylmethoxy)-pyrimidin-4-yloxy]-naphthalen-1-yl}-urea;~~
1-{4-[2-(3-Dimethylamino-propylamino)-6-methyl-pyrimidin-4-yloxy]-naphthalen-1-yl}-3-[2-methoxy-5-(2,2,2-trifluoro-ethyl)-phenyl]-urea;
~~1-[2-Methoxy-5-(2,2,2-trifluoro-ethyl)-phenyl]-3-{4-[6-methyl-2-(3-morpholin-4-yl-propylamino)-pyrimidin-4-yloxy]-naphthalen-1-yl}-urea;~~
~~1-(2-Methoxy-5-trifluoromethoxy-phenyl)-3-{4-[6-methyl-2-(3-morpholin-4-yl-propylamino)-pyrimidin-4-yloxy]-naphthalen-1-yl}-urea;~~
1-[4-(2-Amino-6-methyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-3-(2-methoxy-5-pentafluoroethyl-phenyl)-urea;
1-{4-[2-(Cyclopropylmethyl-amino)-6-methyl-pyrimidin-4-yloxy]-naphthalen-1-yl}-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea;
1-{4-[2-(2-Dimethylamino-ethylamino)-6-methyl-pyrimidin-4-yloxy]-naphthalen-1-yl}-3-[2-methoxy-5-(2,2,2-trifluoro-1-trifluoromethyl-ethyl)-phenyl]-urea;
~~1-[2-Methoxy-5-(2,2,2-trifluoro-1-trifluoromethyl-ethyl)-phenyl]-3-[4-(6-methyl-2-morpholin-4-ylmethyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-urea;~~

~~1-(2-Methoxy-5-trifluoromethyl-phenyl)-3-{4-[6-methyl-2-(4-methyl-piperazin-1-ylmethyl)-pyrimidin-4-yloxy]-naphthalen-1-yl}-urea;~~ and
1-[4-(2-Dimethylaminomethyl-6-methyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea

or the pharmaceutically acceptable acids, esters, salts or isomers thereof.

Claim 9 (currently amended): A compound chosen from

1-[4-(2-Amino-6-methyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea;
1-(2-Methoxy-5-trifluoromethyl-phenyl)-3-[4-(6-methyl-2-methylamino-pyrimidin-4-yloxy)-naphthalen-1-yl]-urea;
~~1-(2-Methoxy-5-trifluoromethyl-phenyl)-3-[4-(6-methyl-2-morpholin-4-ylmethyl-pyrimidin-4-yloxy)-naphthalen-1-yl]-urea;~~
1-{4-[2-(2-Dimethylamino-ethylamino)-6-methyl-pyrimidin-4-yloxy]-naphthalen-1-yl}-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea <u>and</u> ~~;~~
1-{4-[2-(2-Methoxy-ethylamino)-6-methyl-pyrimidin-4-yloxy]-naphthalen-1-yl}-3-(2-methoxy-5-trifluoromethyl-phenyl)-urea;
~~1-(2-Methoxy-5-trifluoromethyl-phenyl)-3-{4-[6-methyl-2-(2-pyrrolidin-1-yl-ethylamino)-pyrimidin-4-yloxy]-naphthalen-1-yl}-urea;~~ ~~and~~
~~1-(2-Methoxy-5-trifluoromethyl-phenyl)-3-{4-[6-methyl-2-(2-morpholin-4-yl-ethylamino)-pyrimidin-4-yloxy]-naphthalen-1-yl}-urea~~

or the pharmaceutically acceptable acids, esters, salts or isomers thereof.

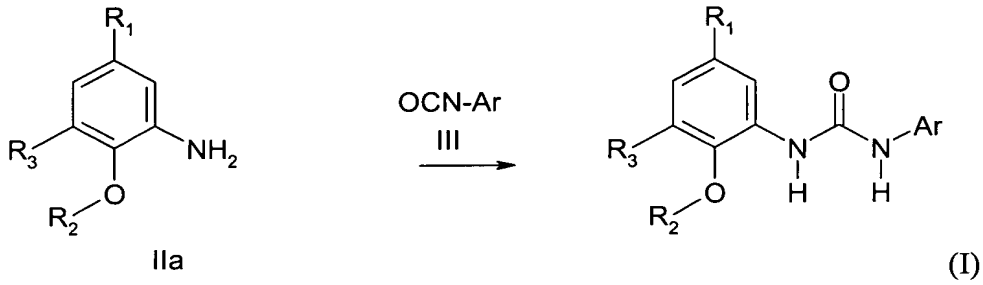
Claim 10 (original): A pharmaceutical composition containing a pharmaceutically effective amount of a compound according to claim 1 and one or more pharmaceutically acceptable carriers and/or adjuvants.

Claim 11 (cancelled).

Claim 12 (currently amended): A ~~The~~ method ~~according to claim 11 wherein cytokine mediated~~ of treating a disease or condition ~~is~~ selected from rheumatoid arthritis, inflammatory bowel disease, septic shock, osteoarthritis, Crohn's disease, ulcerative colitis, multiple sclerosis, Guillain-Barre syndrome, psoriasis, graft versus host disease, systemic lupus erythematosus, restenosis following percutaneous transluminal coronary angioplasty, diabetes, toxic shock syndrome, Alzheimer's disease, acute and chronic pain, contact dermatitis, atherosclerosis, traumatic arthritis, glomerulonephritis, reperfusion injury, sepsis, bone resorption diseases, chronic obstructive pulmonary disease, congestive heart failure, asthma, stroke, myocardial infarction, thermal injury, adult respiratory distress syndrome (ARDS), multiple organ injury secondary to trauma, dermatoses with acute inflammatory components, acute purulent meningitis, necrotizing enterocolitis and syndromes associated with hemodialysis, leukopherisis and granulocyte transfusion.

Claim 13 (withdrawn): A process for preparing compounds of the formula (I) according to claim 1, wherein:

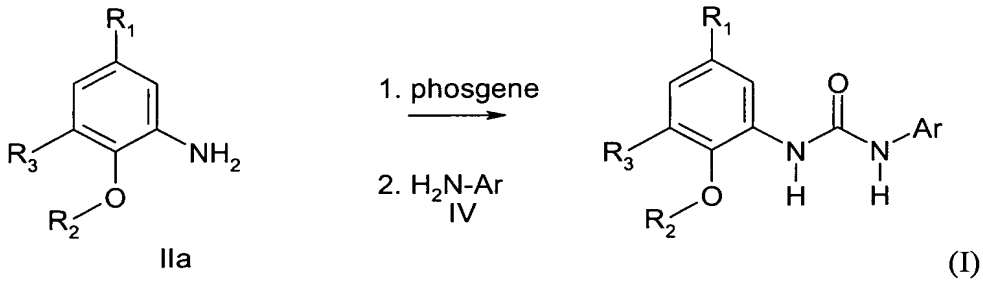
a) an arylamine of formula IIa and an arylisocyanate of formula III is dissolved in a non-protic, anhydrous solvent chosen from THF, ether, toluene, dioxane or ethyl acetate, between 0 - 45° C, for 2-24 h, and subsequently isolating the product of formula I or precursors thereof;



IIa (I)

or

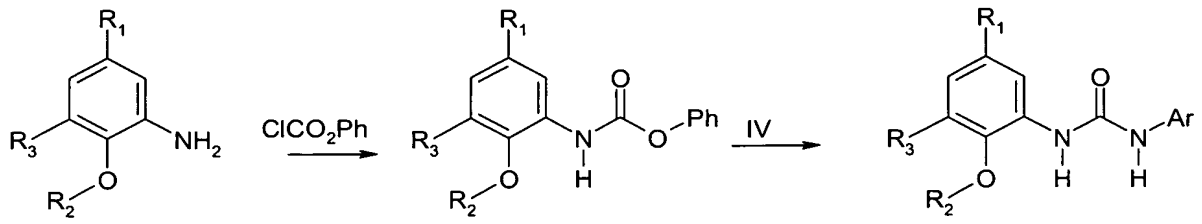
b) dissolving an arylamine of formula IIa in a halogenated solvent, diluting the mixture with aqueous alkali and combining with phosgene to provide the corresponding isocyanate, mixing the isocyanate and arylamine IV in a non-protic, anhydrous solvent selected from THF, ether, toluene, dioxane, dichloromethane or ethyl acetate at between about 0 - 45° C and subsequently isolating to provide the product of formula I or precursors thereof;



IIa (I)

or

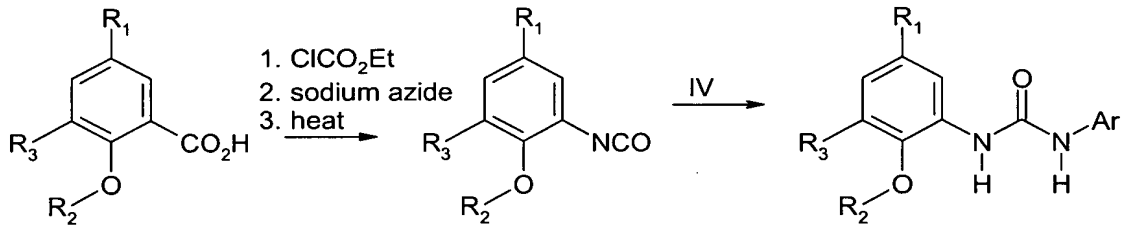
c) dissolving an arylamine of formula IIa in a suitable halogenated solvent optionally adding a suitable base followed by an alkyl or aryl chloroformate between 0 - 85° C, for 2 - 24 h, providing carbamate V, mixing carbamate V and arylamine IV in a non-protic, anhydrous solvent selected from THF, ether, toluene, dioxane, dichloromethane or ethyl acetate, between 0 - 110 ^{0}C, for 2 - 24 h, subsequently isolating to provide the product of formula I or precursors thereof;



IIa V (I)

or

d) dissolving an aromatic carboxylic acid in a non-protic solvent and an inorganic base, cooling the mixture is to –30- 0°C, adding an alkyl chloroformate at 0°C for 1-3 hours, adding sodium azide in water diluted with toluene followed by heating for 1-4 hours, cooling to room temperature to give isocyanate (Va), reacting isocyanate (Va) with amine (IV), subsequently isolating to give product of formula I or precursors thereof;



IIb Va (I).